



## T-cell immunodeficiency, congenital alopecia, and nail dystrophy

T-cell immunodeficiency, congenital alopecia, and nail dystrophy is a type of severe combined immunodeficiency (SCID), which is a group of disorders characterized by an almost total lack of immune protection from foreign invaders such as bacteria and viruses. People with this form of SCID are missing functional immune cells called T cells, which normally recognize and attack foreign invaders to prevent infection. Without functional T cells, affected individuals develop repeated and persistent infections starting early in life. The infections result in slow growth and can be life-threatening; without effective treatment, most affected individuals live only into infancy or early childhood.

T-cell immunodeficiency, congenital alopecia, and nail dystrophy also affects growth of the hair and nails. Congenital alopecia refers to an absence of hair that is apparent from birth. Affected individuals have no scalp hair, eyebrows, or eyelashes. Nail dystrophy is a general term that describes malformed fingernails and toenails; in this condition, the nails are often ridged, pitted, or abnormally curved.

Researchers have described abnormalities of the brain and spinal cord (central nervous system) in at least two cases of this condition. However, it is not yet known whether central nervous system abnormalities are a common feature of T-cell immunodeficiency, congenital alopecia, and nail dystrophy.

### Frequency

T-cell immunodeficiency, congenital alopecia, and nail dystrophy is a rare disorder. It has been diagnosed in only a few individuals, almost all of whom are members of a large extended family from a community in southern Italy.

### Genetic Changes

T-cell immunodeficiency, congenital alopecia, and nail dystrophy results from mutations in the *FOXN1* gene. This gene provides instructions for making a protein that is important for development of the skin, hair, nails, and immune system. Studies suggest that this protein helps guide the formation of hair follicles and the growth of fingernails and toenails. The FOXN1 protein also plays a critical role in the formation of the thymus, which is a gland located behind the breastbone where T cells mature and become functional. Researchers suspect that the FOXN1 protein is also involved in the development of the central nervous system, although its role is unclear.

Mutations in the *FOXN1* gene prevent cells from making any functional FOXN1 protein. Without this protein, hair and nails cannot grow normally. A lack of FOXN1 protein also prevents the formation of the thymus. When this gland is not present, the immune system cannot produce mature, functional T cells to fight infections. As a result, people with T-cell immunodeficiency, congenital alopecia, and nail dystrophy develop recurrent serious infections starting early in life.

## **Inheritance Pattern**

This condition is inherited in an autosomal recessive pattern, which means both copies of the gene in each cell have mutations. The parents of an individual with an autosomal recessive condition each carry one copy of the mutated gene, but they typically do not show signs and symptoms of the condition. However, some people who carry one copy of a mutated *FOXN1* gene have abnormal fingernails or toenails.

## **Other Names for This Condition**

- alymphoid cystic thymic dysgenesis
- congenital alopecia and nail dystrophy associated with severe functional T-cell immunodeficiency
- Pignata Guarino syndrome
- winged helix deficiency

## **Diagnosis & Management**

These resources address the diagnosis or management of T-cell immunodeficiency, congenital alopecia, and nail dystrophy:

- Be The Match: What is a Bone Marrow Transplant?  
<https://bethematch.org/for-patients-and-families/considering-transplant-and-other-treatment-options/what-is-a-bone-marrow-transplant/>
- Genetic Testing Registry: T-cell immunodeficiency, congenital alopecia and nail dystrophy  
<https://www.ncbi.nlm.nih.gov/gtr/conditions/C1866426/>
- MedlinePlus Encyclopedia: Bone Marrow Transplant  
<https://medlineplus.gov/ency/article/003009.htm>

These resources from MedlinePlus offer information about the diagnosis and management of various health conditions:

- Diagnostic Tests  
<https://medlineplus.gov/diagnostictests.html>
- Drug Therapy  
<https://medlineplus.gov/drugtherapy.html>

- Surgery and Rehabilitation  
<https://medlineplus.gov/surgeryandrehabilitation.html>
- Genetic Counseling  
<https://medlineplus.gov/geneticcounseling.html>
- Palliative Care  
<https://medlineplus.gov/palliativecare.html>

## **Additional Information & Resources**

### MedlinePlus

- Encyclopedia: Bone Marrow Transplant  
<https://medlineplus.gov/ency/article/003009.htm>
- Encyclopedia: Immune Response (Video)  
<https://medlineplus.gov/ency/anatomyvideos/000073.htm>
- Encyclopedia: Immunodeficiency Disorders  
<https://medlineplus.gov/ency/article/000818.htm>
- Health Topic: Hair Loss  
<https://medlineplus.gov/hairloss.html>
- Health Topic: Immune System and Disorders  
<https://medlineplus.gov/immunesystemanddisorders.html>
- Health Topic: Nail Diseases  
<https://medlineplus.gov/naildiseases.html>

### Genetic and Rare Diseases Information Center

- T-cell immunodeficiency, congenital alopecia and nail dystrophy  
<https://rarediseases.info.nih.gov/diseases/4358/t-cell-immunodeficiency-congenital-alopecia-and-nail-dystrophy>

### Additional NIH Resources

- National Institute of Allergy and Infectious Diseases: Primary Immune Deficiency Diseases  
<https://www.niaid.nih.gov/diseases-conditions/primary-immune-deficiency-diseases-pids>

## Educational Resources

- Boston Children's Hospital: Severe Combined Immunodeficiency  
<http://www.childrenshospital.org/conditions-and-treatments/conditions/severe-combined-immunodeficiency-scid>
- Disease InfoSearch: T-cell immunodeficiency, congenital alopecia and nail dystrophy  
<http://www.diseaseinfosearch.org/T-cell+immunodeficiency%2C+congenital+alopecia+and+nail+dystrophy/6987>
- Great Ormond Street Hospital for Children: Severe Combined Immunodeficiency  
<http://www.gosh.nhs.uk/medical-information-0/search-medical-conditions/severe-combined-immunodeficiency-scid>
- KidsHealth from Nemours: Severe Combined Immunodeficiency  
<http://kidshealth.org/en/parents/severe-immunodeficiency.html>
- MalaCards: t-cell immunodeficiency, congenital alopecia, and nail dystrophy  
[http://www.malacards.org/card/t\\_cell\\_immunodeficiency\\_congenital\\_alopecia\\_and\\_nail\\_dystrophy\\_2](http://www.malacards.org/card/t_cell_immunodeficiency_congenital_alopecia_and_nail_dystrophy_2)
- Merck Manual Consumer Version: Hair Loss (Alopecia)  
<http://www.merckmanuals.com/home/skin-disorders/hair-disorders/alopecia-hair-loss>
- Orphanet: A lymphoid cystic thymic dysgenesis  
[http://www.orpha.net/consor/cgi-bin/OC\\_Exp.php?Lng=EN&Expert=169095](http://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=169095)

## Patient Support and Advocacy Resources

- Immune Deficiency Foundation  
<http://primaryimmune.org/>
- International Patient Organisation for Primary Immunodeficiencies  
<http://www.ipopi.org/>
- Jeffrey Modell Foundation  
<http://www.info4pi.org/>
- National Organization for Rare Disorders (NORD): Severe Combined Immunodeficiency  
<https://rarediseases.org/rare-diseases/severe-combined-immunodeficiency/>
- Resource List from the University of Kansas Medical Center: Immune Deficiency Conditions  
<http://www.kumc.edu/gec/support/immune.html>

## Genetic Testing Registry

- T-cell immunodeficiency, congenital alopecia and nail dystrophy  
<https://www.ncbi.nlm.nih.gov/gtr/conditions/C1866426/>

## Scientific Articles on PubMed

- PubMed  
<https://www.ncbi.nlm.nih.gov/pubmed?term=%28t-cell+immunodeficiency,+congenital+alopecia,+and+nail+dystrophy%29+OR+%28%28alopecia%5BTIAB%5D%29+AND+%28immunodeficiency%5BTIAB%5D%29+AND+%28nail+dystrophy%5BTIAB%5D%29%29+OR+%28%28immunodeficiency%5BTIAB%5D%29+AND+%28FOXN1%5BTIAB%5D%29%29+OR+%28%28T-cell%5BTI%5D%29+AND+%28FOXN1%5BTI%5D%29%29+NOT+%28zinc%5BTI%5D%29+AND+english%5BIa%5D+AND+human%5Bmh%5D>

## OMIM

- T-CELL IMMUNODEFICIENCY, CONGENITAL ALOPECIA, AND NAIL DYSTROPHY  
<http://omim.org/entry/601705>

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